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**Patent and Trademark Office**

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APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.
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09/684,725    10/06/00    HARLAND

L    PCS10361ADAM

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HM12/1102

EXAMINER
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GREGG C BENSON  
PFIZER INC  
PATENT DEPARTMENT MS  
4159 EASTERN POINT ROAD  
GROTON CT 06340

L I R	
ART UNIT	PAPER NUMBER

1646  
DATE MAILED:

11/02/01

8

**Please find below and/or attached an Office communication concerning this application or proceeding.**

**Commissioner of Patents and Trademarks**

# Office Action Summary

Application No.

09/684,725

Applicant(s)

HARLAND, LEE

Examiner

Ruixiang Li

Art Unit

1646

-- The MAILING DATE of this communication appears on the cover sheet with the correspondence address --

## Period for Reply

A SHORTENED STATUTORY PERIOD FOR REPLY IS SET TO EXPIRE 3 MONTH(S) FROM THE MAILING DATE OF THIS COMMUNICATION.

- Extensions of time may be available under the provisions of 37 CFR 1.136(a). In no event, however, may a reply be timely filed after SIX (6) MONTHS from the mailing date of this communication.
- If the period for reply specified above is less than thirty (30) days, a reply within the statutory minimum of thirty (30) days will be considered timely.
- If NO period for reply is specified above, the maximum statutory period will apply and will expire SIX (6) MONTHS from the mailing date of this communication.
- Failure to reply within the set or extended period for reply will, by statute, cause the application to become ABANDONED (35 U.S.C. § 133).
- Any reply received by the Office later than three months after the mailing date of this communication, even if timely filed, may reduce any earned patent term adjustment. See 37 CFR 1.704(b).

## Status

- 1) ☐ Responsive to communication(s) filed on \_\_\_\_.
- 2a) ☐ This action is FINAL. 2b) ☒ This action is non-final.
- 3) ☐ Since this application is in condition for allowance except for formal matters, prosecution as to the merits is closed in accordance with the practice under *Ex parte Quayle*, 1935 C.D. 11, 453 O.G. 213.

## Disposition of Claims

- 4) ☒ Claim(s) 1-12 and 22 is/are pending in the application.
- 4a) Of the above claim(s) 13-21 is/are withdrawn from consideration.
- 5) ☐ Claim(s) \_\_\_\_ is/are allowed.
- 6) ☒ Claim(s) 1-12 and 22 is/are rejected.
- 7) ☐ Claim(s) \_\_\_\_ is/are objected to.
- 8) ☒ Claim(s) 1-22 are subject to restriction and/or election requirement.

## Application Papers

- 9) ☒ The specification is objected to by the Examiner.
- 10) ☒ The drawing(s) filed on 10/06/2000 is/are: a) ☒ accepted or b) ☐ objected to by the Examiner.
- Applicant may not request that any objection to the drawing(s) be held in abeyance. See 37 CFR 1.85(a).
- 11) ☐ The proposed drawing correction filed on \_\_\_\_ is: a) ☐ approved b) ☐ disapproved by the Examiner.
- If approved, corrected drawings are required in reply to this Office action.
- 12) ☐ The oath or declaration is objected to by the Examiner.

## Priority under 35 U.S.C. §§ 119 and 120

- 13) ☒ Acknowledgment is made of a claim for foreign priority under 35 U.S.C. § 119(a)-(d) or (f).
- a) ☐ All b) ☒ Some \* c) ☐ None of:
1. ☒ Certified copies of the priority documents have been received.
2. ☐ Certified copies of the priority documents have been received in Application No. \_\_\_\_.
3. ☐ Copies of the certified copies of the priority documents have been received in this National Stage application from the International Bureau (PCT Rule 17.2(a)).
- \* See the attached detailed Office action for a list of the certified copies not received.
- 14) ☐ Acknowledgment is made of a claim for domestic priority under 35 U.S.C. § 119(e) (to a provisional application).
- a) ☐ The translation of the foreign language provisional application has been received.
- 15) ☐ Acknowledgment is made of a claim for domestic priority under 35 U.S.C. §§ 120 and/or 121.

## Attachment(s)

- 1) ☐ Notice of References Cited (PTO-892)
- 2) ☐ Notice of Draftsperson's Patent Drawing Review (PTO-948)
- 3) ☐ Information Disclosure Statement(s) (PTO-1449) Paper No(s) \_\_\_\_.
- 4) ☐ Interview Summary (PTO-413) Paper No(s). \_\_\_\_.
- 5) ☐ Notice of Informal Patent Application (PTO-152)
- 6) ☐ Other: \_\_\_\_\_.

## **DETAILED ACTION**

### ***Election/Restrictions***

1. Restriction to one of the following inventions is required under 35 U.S.C. 121:

- I. Claims 1-12, and 22, drawn to polynucleotides, primers, vectors, host cells, and the process of making the polypeptide, classified in class 536, subclasses 23.1 and 24.33; class 435, subclasses 320.1, 325, and 69.1.
- II. Claim 13, drawn to polypeptides, classified in class 530, subclass 324.
- III. Claims 14 and 16, drawn to antibodies, classified in class 530, subclass 387.9.
- IV. Claims 15 and 17, drawn to polypeptide modulators, classified in class 530, subclass 300.
- V. Claim 18, drawn to a method of identifying peptide-binding compounds, classified in class 435, subclass 7.1.
- VI. Claim 19, drawn to a method of treatment with antibodies, classified in class 424, subclass 134.1.
- VII. Claims 20 and 21, drawn to a method of treatment with peptide modulators, classified in class 514, subclass 2.

2. The inventions are distinct, each from the other for the following reasons. Inventions

I, II, III, and IV are unrelated. Inventions are unrelated if it can be shown that they are not disclosed as capable of use together and they have different modes of operation, different functions, or different effects (MPEP §806.04, MPEP §808.01).

In the instance case, the different inventions are drawn to completely different

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products having completely different structures and biological functions which are not interchangeable and which require non-cohesive searches and considerations.

3. Inventions V, VI and VII are unrelated. Inventions are unrelated if it can be shown that they are not disclosed as capable of use together and they have different modes of operation, different functions, or different effects (MPEP §806.04, MPEP §808.01). In the instance case the different inventions are drawn to completely different methods each having completely different method steps, using different compositions, and having completely different outcomes. The method of identifying peptide-binding compounds will not provide information regarding the method of treatment with either an antibody or a peptide modulator; On the other hand, the method of treatment with either an antibody or a peptide modulator will not be able to identify the peptide-binding compounds and the three methods are exclusive.
4. Inventions II and V are related as product and process of use. The inventions can be shown to be distinct if either or both of the following can be shown: (1) the process for using the product as claimed can be practiced with another materially different product or (2) the product as claimed can be used in a materially different process of using that product (MPEP § 806.05(h)). In the instant case, the polypeptide may be used in materially different methods, such as production of antibody by immunization of the mice.
5. Inventions III and VI are related as product and process of use. The inventions can be shown to be distinct if either or both of the following can be shown: (1) the process for using the product as claimed can be practiced with another materially

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different product or (2) the product as claimed can be used in a materially different process of using that product (MPEP § 806.05(h)). In the instant case, the antibodies may be used to detect or isolate the peptide.

6. Inventions IV and VII are related as product and process of use. The inventions can be shown to be distinct if either or both of the following can be shown: (1) the process for using the product as claimed can be practiced with another materially different product or (2) the product as claimed can be used in a materially different process of using that product (MPEP § 806.05(h)). In the instant case the peptide modulators can be used in a binding assay or to study the biological functions of the peptide.
7. Invention I is an independent invention from V, VI, and VII; Invention II is an independent invention from VI and VII; Invention III is an independent invention from V and VII; Invention IV is an independent invention from V and VI. The different inventions are drawn to distinct product and method inventions.
8. Because these inventions are distinct for the reasons given above and have acquired a separate status in the art as shown by their different classification, restriction for examination purposes as indicated is proper.
9. Because these inventions are distinct for the reasons given above and have acquired a separate status in the art because of their recognized divergent subject matter, restriction for examination purposes as indicated is proper.

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10. Because these inventions are distinct for the reasons given above and the search required for a single group is not required for any other group, restriction for examination purposes as indicated is proper.
11. During a telephone conversation with Deborah A. Martin on August 9, a provisional election was made without traverse to prosecute the invention of group I, claims 1-12 and 22. Affirmation of this election must be made by applicant in replying to this Office action. Claims 13-21 are withdrawn from further consideration by the examiner, 37 CFR 1.142(b), as being drawn to a non-elected invention.

Applicant is reminded that upon the cancellation of claims to a non-elected invention, the inventorship must be amended in compliance with 37 CFR 1.48 (b) if one or more of the currently named inventors is no longer an inventor of at least one claim remaining in the application. Any amendment of inventorship must be accompanied by a petition under 37 CFR 1.48 (b) and by the fee required under 37 CFR 1.17 (I).

***Minor Objection to the Application***

12. The title of the invention is not descriptive. A new title is required that is clearly indicative of the invention to which the claims are directed. The following title is suggested: "A Subtype of Neuromedin U Receptor".

***Claim Rejections—35 USC § 101***

13. 35 U.S.C. 101 reads as follows:

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Whoever invents or discovers any new and useful process, machine, manufacture, or composition of matter, or any new and useful improvement thereof, may obtain a patent therefor, subject to the conditions and requirements of this title.

14. Claims 1-12 and 22 are rejected under 35 U.S.C. 101 because the claimed invention is not supported by either a specific and substantial asserted utility or a well-established utility.

Claims 1-12 and 22 are drawn to polynucleotides encoding G-protein-coupled receptors and methods of expressing these proteins. The claimed polynucleotides are not supported by either a specific and substantial asserted utility or a well-established utility. A specific and substantial utility is one that is particular to the subject matter claimed and that identifies a "real world" context of use for the claimed invention which does not require further research. The asserted utility of use of compounds (agonists or antagonists of the claimed polypeptide) in the manufacture of a medicament for treatment of obesity (page 8, line 36-page 8, line 11), while specific, is not substantial because it would require or constitute carrying out further research to identify or reasonably confirm a "real world" context of use. See *Brenner v. Manson*, 383 U.S. 519, 148 USPQ 689 (Sup. Ct. 1966), noting that "a patent is not a hunting license. It is not a reward for the search, but compensation for its successful conclusion."

Likewise, the following asserted utility (page 9, lines 15-22) is not specific and substantial, "Drugs that modulate the novel PFI-002 receptor will therefore be likely to modulate signal transduction processes. It is therefore likely that modulators of the PFI-002 receptor may be useful for the treatment of many different disorders associated with signal transduction that will most likely include,

but not limited to, obesity, diabetes...". The word "likely" and the disclosure of a list of disorders in the specification clearly indicate that the asserted utility requires further research to determine the specific role of the claimed novel PFI-002 receptor in signal transduction processes and its association with a specific disorder.

In addition, the asserted utility of the PFI-002 polypeptide and nucleotide sequences encoding the polypeptide for screening drug candidates for treatment of diseases (page 9, lines 24-35) is not specific and substantial, either. Furthermore, the specification asserts that nucleotide sequences encoding a PFI-002 receptor provide probes to detect chromosomal aberration (page 19, lines 15-21). Since a disease specifically associated with an abnormal level of nucleotide sequences encoding a PFI-002 receptor has not been identified, the use of the probe to diagnose such chromosomal aberration would require further research. Thus, the asserted utility is not specific and substantial.

The invention also lacks a well-established utility. A well-established utility is a specific, substantial, and creditable utility that is well known, immediately apparent, or implied by the specification's disclosure of the properties of a material. Neither the specification as filed nor any art of record before the priority date of this application discloses or suggests any property or activity for the PFI-002 polypeptide and/or nucleotide sequences encoding the polypeptide such that another non-asserted utility would be well-established for the compounds.

15. Claims 1-12 and 22 are also rejected under 35 U.S.C. 112, first paragraph. Specifically, since the claimed invention is not supported by either a specific and



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substantial asserted utility or a well established utility for the reasons set forth above, one skilled in the art clearly would not know how to use the claimed invention.

Further, even if the polynucleotide of SEQ ID NO: 1 or encoding a polypeptide of SEQ ID NO: 2 were to have a patentable use, the instant disclosure would not be found to be enabling for the claimed genus of homologues or fragments of the polynucleotides.

The factors to be considered when determining whether a disclosure satisfies enablement requirement include: (i) the quantity of experimentation necessary; (ii) the amount of direction or guidance presented; (iii) the existence of working examples; (iv) the nature of the invention; (v) the state of the prior art; (vi) the relative skill of those in the art; (vii) the predictability or unpredictability of the art; and (viii) the breadth of the claims. *Ex Parte Forman*, 230 USPQ 546 (Bd Pat. App. & Int. 1986); *In re Wands*, 858 F. 2d 731, 8 USPQ 2d 1400 (Fed. Cir. 1988).

Each of the claims recites a genus of polynucleotides that has at least 70% (75%, 80%, 85%, 90%, or 95%) identity to a polynucleotide comprising SEQ ID NO: 1 or encoding a polypeptide of SEQ ID NO: 2, as well as polynucleotides comprising fragments of a polynucleotide of SEQ ID NO: 1 or encoding SEQ ID NO: 2. However, other than SEQ ID NOS: 1 and 2, the disclosure does not provide sufficient guidance and information regarding the structural and functional requirements commensurate in scope with what is encompassed by the instant claims. The disclosure has not shown what modifications (e.g., substitutions,

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deletions or additions) one can make to SEQ ID NO: 1 or SEQ ID NO: 2 will result in mutants with the same function as SEQ ID NO: 1 or SEQ ID NO: 2. The state of the art (See, e.g., Ngo, et al, *The Protein Folding Problem and Tertiary Structure Prediction*, 1994, Merz, et al. (ed.), Birkhauser, Boston, MA, pp. 433 and 492-495) is such that the sequence of a peptide and its activity is not well understood and is not predictable. Excising out portions of a protein or modifications to a protein, e.g., by substitutions and deletions, would often result in deleterious effects to the overall activity and effectiveness of the protein.

The instant claims also embrace polynucleotide fragments of any size. However, the disclosure has not shown which portions or fragments of SEQ ID NO: 1 or SEQ ID NO: 2 are critical to the activity of the polypeptide encoded by the claimed polynucleotides. Thus, the disclosure has not provided sufficient guidance and information to enable one skilled in the art to predict which if any fragments of the whole molecule would be reasonably expected to retain characteristic activities alone. The general disclosure that one could make and use SEQ ID NO: 1 or SEQ ID NO: 2 could not be used to be such guidance as to guide one of skill in the art to make and use the invention commensurate in scope with the claims.

Accordingly, the disclosure fails to enable such a myriad of the claimed polynucleotides that not only vary substantially in length but also in amino acid composition and to provide any guidance to those skilled generally on how to make and use useful polynucleotides. Thus, it would require undue experimentation for one skilled in the art to make and use the polynucleotides embraced by the instant

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claims.

In addition, the disclosure provides the asserted use of four probes represented by SEQ ID NOS 3-6 in isolation of PFI-002 and determination of tissue distribution of PFI-002. However, since Claim 8, as it is written, comprises at least 15 contiguous nucleotides, it encompasses virtually any random sequence of any length as long as it has a stretch of at least 15 consecutive nucleotides that is the same as in the sequence(s) recited in claim 1. The state of the art is such that determining the specificity of hybridization probes is empirical by nature and the effect of mismatches within an oligonucleotide probe or primer is unpredictable. In view of the factors, the empirical and unpredictable nature of the art and the lack of guidance with respect to how to use other probes within the scope of the claim, the disclosure does not teach one skilled in the art how to successfully use probes of the claimed scope without undue experimentation.

***Claim Rejections—35 USC § 112, 1<sup>st</sup> paragraph***

16. The following is a quotation of the first paragraph of 35 U.S.C. 112:

The specification shall contain a written description of the invention, and of the manner and process of making and using it, in such full, clear, concise, and exact terms as to enable any person skilled in the art to which it pertains, or with which it is most nearly connected, to make and use the same and shall set forth the best mode contemplated by the inventor of carrying out his invention.

17. Claims 1-12 and 22 are rejected under 35 U.S.C. 112, first paragraph, as containing subject matter which was not described in the specification in such a way as to reasonably convey to one skilled in the relevant art that the inventor, at the time the application was filed, had possession of the claimed invention.

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The description discloses a nucleotide sequence set forth in SEQ ID NO: 1, which encodes a polypeptide as set in SEQ ID NO: 2. However, each of these claims as written includes a genus of polynucleotides that has at least 70% (75%, 80%, 85%, 90%, or 95%) identity to a polynucleotide comprising SEQ ID NO 1 or encoding SEQ ID NO 2, or a polynucleotide encoding the polypeptide expressed by the DNA contained in the clone deposited as NCIMB 41066. Thus, the claims encompass a huge number of polynucleotides that vary substantially both in length and in amino acid composition.

The instant disclosure of a single species of nucleic acid of SEQ ID NO: 1 encoding a single polypeptide of SEQ ID NO: 2 does not adequately support the scope of the claimed genus, which encompasses a substantial variety of subgenera including full-length genes. A description of a genus of cDNA may be achieved by means of a recitation of a representative number of cDNAs, defined by nucleotide sequence, falling within the scope of the genus, or of a recitation of structural features common to the genus, which features constitute a substantial portion of the genus. *Regents of the University of California v. Eli Lilly & Co.*, 119 F3d 1559, 1569, 43 USPQ2d 1398, 1406 (Fed. Cir. 1997). The instant disclosure fails to provide sufficient description information, such as definitive structural or functional features of the claimed genus of polynucleotides. There is no description of the conserved regions that are critical to the structure and function of the genus claimed. There is no description of the sites at which variability may be tolerated and there is no information regarding the relation of structure to function. Furthermore, the prior art

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does not provide compensatory structural or correlative teachings to enable one skilled in the art to identify the encompassed polynucleotides as being identical to those instantly claimed.

Due to the breadth of the claim genus and lack of the definitive structural or functional features of the claimed genus, one skilled in the art would not recognize from the disclosure that the applicant was in possession of the claimed genus.

***Claim Rejections—35 USC § 112, 2<sup>nd</sup> paragraph***

18. The following is a quotation of the second paragraph of 35 U.S.C. 112:

The specification shall conclude with one or more claims particularly pointing out and distinctly claiming the subject matter which the applicant regards as his invention.

19. Claims 1, 8, and 22 are rejected under 35 U.S.C. 112, second paragraph, as being indefinite for failing to particularly point out and distinctly claim the subject matter which applicant regards as the invention.

Claim 1 is indefinite because it recites an abbreviated term, "NCIMB", which should be spelled out in the claim. In addition, the claim can be more clearly stated as "a polynucleotide encoding the polypeptide expressed by the DNA contained in the clone deposited as National Collections of Industrial and Marine Bacteria Limited (NCIMB) 41066";

Claim 8 is vague and indefinite because it is not clear which polynucleotide of claim 1 is referred to by "said polynucleotide".

Claim 22 is indefinite because it uses the words "and/or" to describe an animal cell genetically modified to increase or decrease the expression of a polynucleotide sequence.

20. Claim 22 is also objected to under 37 CFR 1.75(c), as being of improper dependent form for failing to further limit the subject matter of a previous claim. Applicant is required to cancel the claim(s), or amend the claim(s) to place the claim(s) in proper dependent form, or rewrite the claim(s) in independent form. Claim 22 depends on an unelected claim (Claim 13). For prosecution purpose, the examiner considers as if it depends upon Claim 1.

***Claim Rejections—35 USC § 102***

21. The following is a quotation of the appropriate paragraphs of 35 U.S.C. 102 that form the basis for the rejections under this section made in this Office action:

A person shall be entitled to a patent unless –

(b) the invention was patented or described in a printed publication in this or a foreign country or in public use or on sale in this country, more than one year prior to the date of application for patent in the United States.

22. Claims 1-10 and 22 are rejected under 35 U.S.C. 102(b) as being anticipated by Database EMBL Accession No. AC008571 (August 3, 1999), which discloses a nucleotide sequence comprising SEQ ID NO: 1 (See attached sequence alignment) as well as a nucleotide sequence capable of hybridizing to the nucleotide sequence. The nucleotide sequence is present in a vector which in turn is present in a host cell, meeting the limitation of claims 9, 10, and 22.

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23. Claim 1 is also rejected under 35 U.S.C. 102(b) as being anticipated by Lubert

Stryer (*Biochemistry*, 3<sup>rd</sup> Edition, pp71-90, W. H. Freeman, 1988). Stryer teaches the structure of nucleotides, DNA, and RNA. Claim 1 recites "a polynucleotide fragment" which encompasses just about everything, from a single nucleotide to at least SEQ ID NO: 1 or its complement. Thus, the reference by Stryer anticipates Claim 1.

24. Claims 1, 7-12, and 22 are rejected under 35 U.S.C. 102(b) as being anticipated by

Tan, et al (IDS, paper #7). Tan et al disclose a nucleotide sequence of a human G-protein-coupled receptor (FM3) with two regions that contain 19 and 26 contiguous nucleotides identical to that set forth in SEQ ID NO: 1 (See attached sequence alignment). Also disclosed are a nucleotide sequence of mouse G-protein-coupled receptor (FM3) with 15 contiguous nucleotides identical to that set forth in SEQ ID NO: 1 (See attached sequence alignment) and the use of a radiolabeled probe encompassing mouse FM-3 ORF in determining the expression profile of FM3 mRNA in several mouse tissues (See Fig. 3, page 228). Furthermore, Tan, et al disclose transfected HEK-293 cells expressing FM3 (cell membrane expression of FM3; See page 228, column 1, lines 24-27), which meets the limitation of claims 10-12 and 22.

25. Claim 12 is also rejected under 35 U.S.C. 102(b) as being anticipated by Maniatis et

al (*Molecular Cloning: A laboratory Manual*. 2<sup>nd</sup> Edition, Book 3, pp17.37-17.41, Cold Spring Harbor Laboratory Press, 1989). Maniatis et al. teach membrane preparation of a cell by centrifugation and thus the reference meets the limitation of

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claim 12, which encompasses membrane preparations indistinguishable from those of the art.

Any inquiry concerning this communication or earlier communications from the examiner should be directed to Ruixiang Li whose telephone number is (703) 306-0282. The examiner can normally be reached on Monday through Friday from 8:30 am to 5:00 pm.

If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Yvonne Eyler, can be reached on (703) 308-6564. The fax phone number for this Group is (703) 305-3014 or (703) 308-4242.

Communications via Internet e-mail regarding this application, other than those under 35 U.S.C. 132 or which otherwise require a signature, may be used by the applicant and should be addressed to [yvonne.eyler@uspto.gov](mailto:yvonne.eyler@uspto.gov).

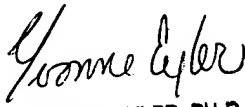
All Internet e-mail communications will be made of record in the application file. PTO employees do not engage in Internet communications where there exists a possibility that sensitive information could be identified or exchanged unless the record includes a properly signed express waiver of the confidentiality requirements of 35 U.S.C. 122. This is more clearly set forth in the Interim Internet Usage Policy published in the Official Gazette of the Patent and Trademark on February 25, 1997 at 1195 OG 89.



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Any inquiry of a general nature or relating to the status of this application or proceeding should be directed to the Group receptionist whose telephone number is (703) 308-0196.

Ruixiang Li  
Examiner  
October 16, 2001

  
YVONNE EYLER, PH.D  
SUPERVISORY PATENT EXAMINER  
TECHNOLOGY CENTER 1600

9	280.8	38.5	1318	89	AF272362	AF272362 Homo sapi
10	280.8	38.5	187451	64	AC017104	AC017104 Homo sapi
11	280.8	35.0	1209	94	AF242873	AF242873 Rattus no
12	255	35.0	1239	94	AB038649	AB038649 Rattus no
13	243	33.3	1218	94	AF044602	AF044602 Mus muscu
14	159.6	21.9	75950	74	AC073449	AC073449 Homo sapi
15	134.8	18.5	2040	88	AF034462	AF034462 Homo sapi
16	134.8	18.5	163284	89	AL137000	AL137000 Homo sapi
17	133.2	18.3	1676	85	AF082210	AF082210 Spheeroid
18	127	17.4	1254	48	E11480	E11480 CDNA encodi
19	127	17.4	4131	93	HSNEURA	X70070 H.sapiens
20	120.4	16.5	870	97	HS060181	U60181 Human growt
21	120.4	16.5	1101	97	HS060179	U60179 Human growt
22	120.4	16.5	146442	74	AC069523	AC069523 Homo sapi
23	119.8	16.4	121652	60	AC008191	AC008191 Drosophi1
24	119.8	16.4	140838	65	AC018176	AC018176 Drosophi1
25	119.8	16.4	219832	60	AC007441	AC007441 Drosophi1
26	119.8	16.4	225374	5	AE003703	AE003703 Drosophi1
27	118.8	16.3	150566	64	AC016938	AC016938 Homo sapi
28	114	15.6	870	7	SS060180	U60180 Sus scrofa
29	114	15.6	1101	7	SS060178	U60178 Sus scrofa
30	112.4	15.4	1161	10	I15508	I15508 Sequence 2
31	112.4	15.4	1161	10	I15508	I15508 Sequence 2
32	112.4	15.4	1370	10	I20930	I20930 Sequence 1
33	112.4	15.4	1370	10	I20930	I20930 Sequence 1
34	112.4	15.4	1466	10	I28195	I28195 Sequence 17
35	112.4	15.4	1466	10	I28195	I28195 Sequence 17
36	112.4	15.4	1504	97	H0MD4C	I28196 Sequence 3
37	112.4	15.4	1504	97	H0MD4C	I28196 Sequence 3
38	112.4	15.4	1610	10	I20932	I20932 Sequence 19
39	110	15.1	1528	95	I28197	I28197 Sequence 5
40	109.2	15.0	1350	94	AB001962	X97121 R.norvegicu
41	109.2	15.0	3129	94	AB001962	AB001962 Rattus no
42	106.6	14.6	3129	94	AB001962	AB001962 Rattus no
43	101.6	13.9	2379	7	BT2DOR	U94321 Rattus norv
44	101	13.9	1089	94	AF149717	X51657 B.taurus do
45	101	13.9	2351	94	AB015645	AF149717 Rattus no
						AB015645 Rattus no

ALIGNMENTS

RESULT 1	AC008571/c	216673 bp	DNA	HTG	05-MAY-2000
LOCUS	Homo sapiens chromosome 5 clone CTC-550M4, WORKING DRAFT SEQUENCE,				
DEFINITION	9 unordered pieces.				
AC008571					
AC008571.3	GI:7708957				
KEYWORDS	HTG; HTGS_PHASE1; HTGS_DRAFT.				
SOURCE	human.				
ORGANISM	Homo sapiens				
	Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;				
	Mammalia; Eutheria; Primates; Catarrhini; Homnidae; Homo.				

REFERENCE 1 (bases 1 to 216673)  
Mammalia; Eutheria; Primates; Catarrhini; Homnidae; Homo.  
DOE Joint Genome Institute.  
Sequencing of Human Chromosome 5  
Unpublished  
2 (bases 1 to 216673)  
DOE Joint Genome Institute.  
Direct Submission  
Submitted (03-ANG-1999) Production Sequencing Facility, DOE Joint  
Genome Institute, 2800 Mitchell Drive, Walnut Creek, CA 94598, USA  
On May 5, 2000 this sequence version replaced gi:7211884.  
-----Genome Center  
Center: Joint Genome Institute  
Center Code: JGI  
Web site: http://www.jgi.doe.gov

Project Information  
Center Project Name: 396672, H361  
Center clone name: CIT-HSPC\_550M4

Summary Statistics	
Consensus quality: 197198 bases at least Q40	
Consensus quality: 207925 bases at least Q30	
Consensus quality: 210803 bases at least Q20	
Estimated insert size: 223000; pulse field gel estimation	
Estimated insert size: 215873; sun-of-coverage estimation	
Quality coverage: 5.46 in Q20 bases; pulse field gel estimation	
Quality coverage: 5.64 in Q20 bases; sun-of-coverage estimation	
* NOTE: This is a 'working draft' sequence. It currently	
* consists of 9 contigs. The true order of the pieces	
* is not known and their order in this sequence record is	
* arbitrary. Gaps between the contigs are represented as	
* runs of N, but the exact sizes of the gaps are unknown.	
* This record will be updated with the finished sequence	
* as soon as it is available and the accession number will	
* be preserved.	
1	3048: contig of 3048 bp in length
3049	3148: gap of unknown length
3149	8405: contig of 5237 bp in length
8406	8505: gap of unknown length
8506	23492: contig of 14987 bp in length
23493	23592: gap of unknown length
40019	40018: contig of 16426 bp in length
40119	40118: gap of unknown length
54777	54776: contig of 14658 bp in length
54877	54876: gap of unknown length
80051	80050: contig of 25174 bp in length
80150	80150: gap of unknown length
121105	121104: contig of 40354 bp in length
121205	121204: gap of unknown length
167091	167091: contig of 45887 bp in length
167092	167091: gap of unknown length
167192	216673: contig of 49482 bp in length.
Location/Qualifiers	
source	1. 216673
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	/db_xref="taxon:9606"
	/clone="CTC-550M4"
	/clone.lib="Caltech human BAC library C"

BASE COUNT 66753 a 41883 c 42226 g 65000 t 811 others

Query Match 100.0%; Score 729; DB 60; Length 216673;  
Best Local Similarity 100.0%; Pred. No. 4e-130;  
Matches 729; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY	1	atggaanaacttcagaatccttcctggaatcaccagacagaagaataagatccatccag	60
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QY	61	aaacacctgaacagacacacagagatattcggccttccttcggagactggcgagacac	120
DB	84343	AAACACCTGAACAGACACGAGAGATATCGGCTTCCTTCGCGAGCTCGGCGACACAC	84284
QY	121	ttcttcctcccgctgctggtatgtatgtacaaatttcgtggtgggtcatatggcaat	180
DB	84283	TTCCTCCTCCCGCTGCTGCTGATGTCAGCAATTTTGTGTGGGTGATGTCGAAT	84224
QY	181	gtccctggtgctggtgattcctgcagacacagagctagaagaagcccaactactac	240
DB	84223	GTCCTGCTGCTGCTGCTGATGTCAGCAACAGGCTATGAAAGCGCCCAACTACTAC	84164
QY	241	ctcttcagctgagctgctcctgacccctcggtgtctctctcttggaatgccccgagtc	300
DB	84163	CTCTTCAGCTGAGCTGCTCCTGACCTCGGTGTCTCTCTTCGAAATGCCCTGAGGTC	84104
QY	301	tatgagatgtgacgaactaccccttctgttcgggcccgtggcgtcactcaagagc	360
DB	84103	TATGAGATGTGACGAACACTACCTTCTGTTCGGGCCCGTGGCGCTGCTACTTCAAGAGC	84044
QY	361	gccctcttgagacgctgtgcttcgctccatcctcagatcaacacacgctcagcgtgag	420

DB 84043 GCCCTCTTTAGACCGGTGCTCTTCCCTCCATCCATGACACACCGCTCAGCGTGAG 83984  
QY 421 CGTCACTGTCACCTCCTACACCCGCTCCGCGCAACCTGAGAGACCCGCGCGG 480  
DB 83983 CGCTACGTGCGCATCTACACCGCTTCCGCGCAACCTGAGACACCGCGCGG 83924  
QY 481 CTGAGATCTCTGCGCTCTGCGGCTCTCTGCTCTCTCTCTCTCTCTCTCTCT 540  
DB 83923 CTGAGATCTCTGCGCATCTCTGCGGCTTCTCTGCTCTCTCTCTCTCTCTCT 83864  
QY 541 ATCAATGATCAAGTCCACCTCTCTCTCTCTCTCTCTCTCTCTCTCTCTCTCTCT 600  
DB 83863 ATCAATGATCAAGTCCACCTCTCTCTCTCTCTCTCTCTCTCTCTCTCTCTCTCT 83804  
QY 601 TGTACGTCATCAAGCCCATGTGATCTCAATTTTCATCATCGATCGATCGATCG 83803  
661 TTCTACT 720  
DB 83743 TTCTACT 83684  
QY 721 AGTATCTAG 729  
DB 83683 AGTATCTAG 83675

## RESULT 2

AF242874 1239 bp mRNA PRI 13-JUL-2000  
LOCUS Homo sapiens neuromedin U receptor 2 (NMU2R) mRNA, complete cds.  
DEFINITION AF242874  
ACCESSION AF242874  
VERSION AF242874.1 GI:9082155  
KEYWORDS

## SOURCE

ORGANISM human.  
Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
Mammalia; Eutheria; Primates; Catarrhini; Homiidae; Homo.

## REFERENCE

## AUTHORS

Howard, A.D., Wang, R., Pong, S.S., Mellin, T.N., Strack, A., Guan, X.M.,  
Zeng, Z., Williams, D.L., Jr., Feigman, S.D., Nunes, C.N., Murphy, B.,  
Stall, J.N., Yu, H., Jiang, Q., Clements, M.K., Tan, C.P., McKee, K.K.,  
Hreniuk, D.L., McDonald, T.P., Lynch, K.R., Evans, J.F., Austin, C.P.,  
Casky, C.T., Van der Ploeg, L.H. and Liu, Q.  
Identification of receptors for neuromedin U and its role in  
feeding

## TITLE

Nature 406 (6791), 70-74 (2000)

## JOURNAL

## MEDLINE

## REFERENCE

## AUTHORS

## TITLE

## JOURNAL

Submitted (09-MAR-2000) Pharmacology, Merck Research Labs, West  
Point, PA 19486, USA

## FEATURES

## SOURCE

1.1239 location/Qualifiers  
/organism="Homo sapiens"  
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/protein\_id="AA82755.1"  
/db\_xref="GI:9082155"  
/translation="MEKIQNASWYQKLEDPFOKHLNSTEETYLICGPRSHRPTLP  
VSVVYPIFVGVIGNVLVLOHQAAMKTPNLYTLFSLAVSDLLVILGLPLEYE  
MRNYPFLPGVPCYKFLAFVFCASLSTTVSEYVAIILHFRKLQSTRRA  
LRLIGVMPGVSIVFSPNSIHGIRHPNSLVGSAIVCTVIRPMIYINFIQVTS  
FLYILPMYIVISLYILMALRLKDKSLDEAGNANIQRCKRSVKMLFLVLVFAI

## CDS

## gene

GAP of reverse of: us-09-684-725-1 . Check: 1088 from: 1 to: 729  
 FROMIG of: /home/bobryen/big/US09684725.seq  
 sequence 1, application us/09684725

general information:  
 applicant: lee harland  
 title of invention: novel polypeptide  
 file reference: pcs10361adam

to: version1. check: 7472 from: 192001 to: 218807

Symbol comparison table: /opt/gcg/gcgcore/data/runningdata/nwsgapdna.comp  
 CompCheck: 8760

Gap Weight: 50 Average Match: 10.000  
 Length Weight: 3 Average Mismatch: 0.000

Quality: 7290 Length: 26807  
 Ratio: 10.000 Gaps: 0  
 Percent Similarity: 100.000 Percent Identity: 100.000

Match display thresholds for the alignment(s):

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us-09-684-725-1 x version1 September 12, 2001 10:41

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729 .....ctagatactcact 717
204851 tgaaaaaaagaggagatcgtcgtcctcaagccaccagcctagatactcact 204900
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666 gtagaataagaaagagagtgacctcgtgatagtgaattgtatgataccatcag 617
204951 gtagaataagaaagagagtgacctcgtgatagtgaattgtatgataccatcag 205000
616 gcttgatgacgtacaggtcgtccgaacctcgtggaccagggaccatctgggg 567
205001 gcttgatgacgtacaggtcgtccgaacctcgtggaccagggaccatctgggg 205050
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205101 caaggaagaagccccaagacatgccaagagatcctgaaggcccgccggcgagg 205150
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205451 ccaccacaaaatttggcacatcacccacagacagcgggaggaagaagtgg 205500
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AC 008571

Version AC008571.1

GI : 5686496

(Submitted on August 3, 1999)

OY	421	cgtacgtagccaccctacacccggttcgcggcacaactgcagagcaaccggcgccggcc	480
Dd	532	CGCATGTGGCCATTGTCACCCCTTCCAGACCAGAGCTGAGAGACACCGCGCAGGGGCC	591
OY	481	ctcgaagtctctggagcatcgtcttgaggcttcttcggtctctctccctggccaacacsac	540
Dd	592	CTCAGGATCTCTCACCCCTAGCTGAGCTCTCTGTGGCTCTTTCTTGCCCAATACCAAC	651
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Dd	652	ATCCATGTGGCATCAAGTTCACACTTTTCCCAAGGGGTCTCCGTACCTGCCTCAGCCACC	711
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Dd	712	TGCACAGTACACAAACCATGTGGGTGAATAAATTGATCATCAAGCTAACAGCTTCCCTC	771
OY	661	ttctacctctctccccagacgtgatcatcagttgtctctctctctaccatcattgacatga	720
Dd	772	TTCTACATCTCCCAAGAATGACCCCTCATGCGCTCTCTACTACCTCATGAGGCTGAGCTG	831
OY	721	a 721	
Dd	832	A 832	
RESULT	8		
HSGOPCR1			
LOCUS	HSGOPCR1	828 bp	DNA PRI 22-JUL-1999
DEFINITION	Homo sapiens orphan G protein-coupled receptor gene, first coding		
ACCESSION	AF044600		
VERSION	AF044600.1	GI:2865467	
KEYWORDS	exon.		
SEGMENT	1 of 2		
SOURCE	human.		
ORGANISM	Homo sapiens		
REFERENCE	Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;		
AUTHORS	Mammalia; Eutheria; Primates; Catarrhini; Homnidae; Homo.		
TITLE	1 (bases 1 to 828)		
JOURNAL	Tan,C.P., McKee,K.R., Liu,Q., Palyha,O.C., Feigheimer,S.D.,		
MEDLINE	Hreniuk,D.L., Smith,R.G. and Howard,A.D.		
PUBMED	Cloning and characterization of a human and murine T-cell orphan		
REFERENCE	G-protein-coupled receptor similar to the growth hormone		
AUTHORS	secretagogue and neurotensin receptors		
TITLE	Genomics 52 (2), 223-229 (1998)		
JOURNAL	99000845		
MEDLINE	9782091		
PUBMED	2 (bases 1 to 828)		
REFERENCE	Tan,C.T., McKee,K.K., Palyha,O.C., Feigheimer,S.D., Hreniuk,D.L.,		
AUTHORS	Smith,R.G. and Howard,A.D.		
TITLE	Direct Submission		
JOURNAL	Submitted (16-JUN-1998) Biochemistry and Physiology, Merck and Co.		
FEATURES	Submitted (16-JUN-1998) Biochemistry and Physiology, Merck and Co.		
SOURCE	Location/Qualifiers		
exon	1..828	/organism="Homo sapiens"	
BASE COUNT	<1..828	/db_xref="taxon:9606"	
ORIGIN	134 a 267 c 256 g 171 t	/note="an intron of about 3kb follows this exon"	
Query Match	38.5%; Score 280.8; DB 93; Length 828;		
Best Local Similarity	65.8%; Pred. No. 4.6e-44;		
Matches 408; Conservative	0; Mismatches 212; Indels 0; Gaps 0;		
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Dd	88	GAGGCCAGCAGACAGAGCTGTTCATGCCCATCTGTGCCACATACCTGATCTTGGG	147
OY	163	gtggaggtcatgtgaatgtcctgtgtgctcgtgtgattctgcagacacaggtatgaag	222

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CD5
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/note="G protein-coupled receptor"
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EALIKYLGPOGLEMPICATYLLIYVVGAVAGLCTVILRRKARTPTNYLFSL
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OHNIVTICIEFYGLSANPLYIKLMSRRPFRRTDIALCLQACCHRLPRHSSHSLSR
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BASE COUNT      208 a      441 c      403 g      266 t
ORIGIN

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Query Match	Best Local Similarity	38.5%;	Score 280.8;	DB 89;	Length 1318;
Matches	408;	Conservative	0;	Mismatches 212;	Indels 0;
					Gaps 0;
QY	103	ggacctgagcgagccactcttcctccccgctgctgtgtgtatgtatgtgcacattttgtg	162		
Db	194	GGGCCCCAGCAGACAGAGCTGTTCATGGCCCAATGTGTGCCATACCTCATCTGTCTGTG	253		
OY	163	gtgggggcatctgtggaatgctcctgtgtgtgctgtgtatcttcgacacaggtcatgaag	222		
Db	254	GTGGGCGCTGTGGGCAATAGGGCTGTACCTGTGTGTGCATCTTCGGCACAAGGCCATYGGC	313		
OY	223	acgcccacaactactactactcttcacgctctgcygtctctgaacctctgtcctcctct	282		
Db	314	ACGGCTACCACTACTACTCTCTTAGCCTTGCCGCGTGTGCACCTCTGTCTGTCTGTG	373		
OY	283	ggaaatgccctgtgaggtctatgatagtgtgtgcgaactaacctcttcctgtgtcggccgtg	342		
Db	374	GGCCGTGCCCCCTGGAGCTGTATAGATGTGGCACAACCTTCCCTCTGTGGCGTTGGT	433		
OY	343	ggctgtactactcaagaagcgccctctctttgagacggtgtgtctcgcctcatctcaagatc	402		
Db	434	GGCTGCTAATTTCCGACCCCTACTGTGTGAGATGTGTCTCCCTGAGCTCAGTCAACGTC	493		
OY	403	acacacgcacagcgtgagagcgctacgtgtgcacatctcaacccgcttcggcgcaaacgtgag	462		
Db	494	ACTGCGCTGAGACGTGTGAACGCTATGTGGCCGTGTGTGCACCCACTTCAAGGCCAAGTTCATG	553		
OY	463	agcacccggcgcgcgccctcagaatctctcgtgcactcgtctcgtgtgtcctccgtctcttc	522		
Db	554	GTGACGCGGGGCCCATGTGTGCGCGAGTCTTGGGCGCGCTGTGGGCTTGTGCATGCTGTGC	613		
OY	523	tcacctggccaaacacagatctcatatgtacataagttcaactactctcccaatgggtcctcg	582		
Db	614	TCCCTGCCCCAACACACAGCTGTGCACGGCATCCGGACGCTGCACGTGCCCTTGGCCGGGCCCA	673		
OY	583	gtccacaggttcgcgcacactcgttacggtacaaagccaatgtgatctaacattcatc	642		
Db	674	GTGCGACAGCTCAGCGTGTTCATGCTGTGGTCCGCCACAGGGCGCTCTTCAACATGTAOTG	733		
OY	643	caggtcacctctctctctatcttactcctcccaatgacgtgtcatcagatgtcctctactac	702		
Db	734	CAGACACACGGCGAGCTCTTCTTGTGCTGTGCCATATGGCCATATGAGAGGTCTCTACTCG	793		
OY	703	ctcatgacatccagatgtgag	722		
Db	794	CTCATTTGGGCTGTGCACATGCG	813		
RESULT	10				
LOCUS	AC017104	187451 bp	DNA	HTG	16-MAR-2001
DEFINITION	Homo sapiens chromosome 2 clone RP11-56215, WORKING DRAFT SEQUENCE,				

## A

-308	ACCTGCCTGCCTCAGCTTCCTTCGCGTTGGGATTAAAGCTGCGCACTACCCTGCCCGCCAATTTTATTATTTTCAAGG	-229
-228	TTTGCACTCCGAATACTCTGTAGTTGAAATGCACCTTAGTGGGTGGCAGATGCTTTCTTTCCAGTGCCATGTGACTAAT	-149
-148	CAGTCTCTACAGTGTGATAACTTAGGACAGAGCTGGGATTACCTAAATGACTTCTGGGTATCTCCCCCTTCTATCCTAC	-69
-68	AGACTCCTCCTGCCTCAATTGTTCCATCTTTCTGGAGCGCTCTCCCCAAATGCTTCAAGGAGCCCC	-1
+ 1	CTG GTC TGC AAT ATC AGT GAG TTC AAG TGG CCC TAT CAA CCT GAG GAT CTG AAC CTT ACC	60
1	L V C N I S E F K W P Y Q P E D L N L T	20
	<b>TM-1</b>	
61	GAT GAG GCC CTG AGG CTG AAG TAT TTG GGG CCA CAG CAG ATG AAA CAG TTT GTC CCC ATC	120
21	D E A L R L K Y L G P Q Q M K Q F V P I	40
121	TGT GTC ACG TAC CTG CTG ATC TTC GTG GTG GGC ACT CTG GGC AAC GGG CTG ACC TGC ACC	180
41	C V T Y L L I F V V G T L G N G L T C T	60
	<b>TM-2</b>	
181	GTC ATC CTG CGC AAC AAG ACT ATG CGC ACG CCC ACC AAC TTC TAC CTC TTC AGC CTC GCT	240
61	V I L R N K T M R T P T N F Y L F S L A	80
241	GTG TCC GAT ATG CTG GTG CTC CTG GTG GGC TTG CCT CTG GAG CTT TAT GAG ATG CAG CAA	300
81	V S D M L V L L V G L P L E L Y E M Q Q	100
	<b>TM-3</b>	
301	AAT TAC CCG TTC CAG CTG GGT GCG AGT GCC TGC TAC TTC CGA ATA CTG CTC TTA GAG ACC	360
101	N Y P F Q L G A S A C Y F R I L L L E T	120
361	GTC TGC CTA GCT TCA GTG CTC AAT GTC ACA GCC CTG AGT GTG GAG CGT TAT GTG GCC GTG	420
121	V C L A S V L N V T A L S V E R Y V A V	140
	<b>TM-4</b>	
421	GTG CGC CCA CTC CAA GCC AAG TCT GTG ATG ACA CGG GCC CAT GTG CGC CGC ATG GTG GGG	480
141	V R P L Q A K S V M T R A H V R R M V G	160
481	GCC ATC TGG GTC CTC GCT ACT CTC TTC TCT CTG CCC AAC ACC AGC CTG CAT GGC CTC AGT	540
161	A I W V L A T L F S L P N T S L H G L S	180
541	CAA CTA ACT GTG CCC TGC CGG GGG CCG GTG CCC GAC TCA GCT ATA TGT TCG CTG GTG GGT	600
181	Q L T V P C R G P V P D S A I C S L V G	200
	<b>TM-5</b>	
601	CCC ATG GAC TTC TAC AAG TTG GTG GTA CTG ACT ACC GCA CTG CTC TTC TTC TGT CTG CCC	660
201	P M D F Y K L V V L T T A L L F F C L P	220
661	ATG GTC ACC ATC AGT GTG CTG TAT CTG CTC ATT GGG CTG CGG CTG CGG AGG GAG AGG ATG	720
221	M V T I S V L Y L L I G L R L R R E R M	240
721	TTG CTC CAA GTG GAG GTC AAG GGC AGG AAA ACC GCA GCA ACC CAG GAG ACC TCC CAC AGA	780
241	L L Q V E V K G R K T A A T Q E T S H R	260
	<b>TM-6</b>	
781	AGG ATT CAG CTG CAA GAT AGG GGA CGG AGA CAG GTG ACC AAG ATG CTG TTT GCA CTG GTT	840
261	R I Q L Q D R G R R Q V T K M L F A L V	280
841	GTG GTA TTC GGC ATC TGC TGG GCT CCA TTC CAT GCT GAC CGT ATC ATG TGG AGC CTG GTG	900
281	V V F G I C W A P F H A D R I M W S L V	300
	<b>TM-7</b>	
901	TAT GGA CAC TCA ACG GAA GGC CTG CAC CTG GCC TAC CAG TGT GTC CAC ATT GCC TCT GGC	960
301	Y G H S T E G L H L A Y Q C V H I A S G	320
961	ATC TTC TTC TAT CTC GGC TCA GCA GCC AAC CCG GTG CTC TAC AGC CTC ATG TCT ACT CGC	1020
321	I F F Y L G S A A N P V L Y S L M S T R	340
1021	TTC CGA GAG ACC TTC CTG CAA GCC CTG GGC CTT GGA ACC CAG TGC TGT CAT CGC CGC CAA	1080
341	F R E T F L Q A L G L G T Q C C H R R Q	360
1081	CCC TAT CAT GGC TCC CAT AAC CAC ATC AGG TTG ACC ACA GGC AGC ACC CTG TGT GAC GTG	1140
361	P Y H G S H N H I R L T T G S T L C D V	380
1141	GGC CAC AGG AAC AGC AGG GAC GAA CCT CTG GCT GTG AAT GAG GAT CCA GGG TGT CAG CAA	1200
381	G H R N S R D E P L A V N E D P G C Q Q	400
1201	GAG ACA GAC CCC TCC TGA	1218
401	E T D P S *	406

FIG. 1. DNA and deduced amino acid sequence for mouse (A) and human (B) FM-3. Putative transmembrane  $\alpha$  helices are overlined and numbered from 1 to 7. For the murine form, a noncanonical leucine codon is postulated to serve as the initiator.

## B

1	ATG	GCT	TGC	AAT	GGC	AGT	GCG	GCC	AGG	GGG	CAC	TTT	GAC	CCT	GAG	GAC	TTG	AAC	CTG	ACT	60
1	M	A	C	N	G	S	A	A	R	G	H	F	D	P	E	D	L	N	L	T	20
TM-1																					
61	GAC	GAG	GCA	CTG	AGA	CTC	AAG	TAC	CTG	GGG	CCC	CAG	CAG	ACA	GAG	CTG	TTC	ATG	CCC	ATC	120
21	D	E	A	L	R	L	K	Y	L	G	P	Q	Q	T	E	L	F	M	P	I	40
TM-2																					
121	TGT	GCC	ACA	TAC	CTG	CTG	ATC	TTC	GTG	GTG	GGC	GCT	GTG	GGC	AAT	GGG	CTG	ACC	TGT	CTG	180
41	C	A	T	Y	L	L	I	F	V	V	G	A	V	G	N	G	L	T	C	L	60
TM-3																					
181	GTC	ATC	CTG	CGC	CAC	AAG	GCC	ATG	CGC	ACG	CCT	ACC	AAC	TAC	TAC	CTC	TTC	AGC	CTG	GCC	240
61	V	I	L	R	H	K	A	M	R	T	P	T	N	Y	Y	L	F	S	L	A	80
TM-4																					
241	GTG	TCG	GAC	CTG	CTG	GTG	CTG	CTG	GTG	GGC	CTG	CCC	CTG	GAG	CTC	TAT	GAG	ATG	TGG	CAC	300
81	V	S	D	L	L	V	L	L	V	G	L	L	E	L	Y	E	M	W	H		100
TM-5																					
301	AAC	TAC	CCC	TTC	CTG	CTG	GGC	GTT	GGT	GGC	TGC	TAT	TTC	CGC	ACG	CTA	CTG	TTT	GAG	ATG	360
101	N	Y	P	F	L	L	G	V	G	G	C	Y	F	R	T	L	L	F	E	M	120
TM-6																					
361	GTC	TGC	CTG	GCC	TCA	GTG	CTC	AAC	GTG	ACT	GCC	CTG	AGC	GTG	GAA	CGC	TAT	GTG	GCC	GTG	420
121	V	C	L	A	S	V	L	N	V	T	A	L	S	V	E	R	Y	V	A	V	140
TM-7																					
421	GTG	CAC	CCA	CTC	CAG	GCC	AGG	TCC	ATG	GTG	ACG	CGG	GCC	CAT	GTG	CGC	CGA	GTG	CTT	GGG	480
141	V	H	P	L	Q	A	R	S	M	V	T	R	A	H	V	R	R	V	L	G	160
TM-8																					
481	GCC	GTC	TGG	GGT	CTT	GCC	ATG	CTC	TGC	TCC	CTG	CCC	AAC	ACC	AGC	CTG	CAC	GGC	ATC	CGG	540
161	A	V	W	G	L	A	M	L	C	S	L	P	N	T	S	L	H	G	I	R	180
TM-9																					
541	CAG	CTG	CAC	GTG	CCC	TGC	CGG	GGC	CCA	GTG	CCA	GAC	TCA	GCT	GTT	TGC	ATG	CTG	GTC	CGC	600
181	Q	L	H	V	P	C	R	G	P	V	P	D	S	A	V	C	M	L	V	R	200
TM-10																					
601	CCA	CGG	GCC	CTC	TAC	AAC	ATG	GTA	GTG	CAG	ACC	ACC	GCG	CTG	CTC	TTC	TTC	TGC	CTG	CCC	660
201	P	R	A	L	Y	N	M	V	V	Q	T	T	A	L	L	F	F	C	L	P	220
TM-11																					
661	ATG	GCC	ATC	ATG	AGC	GTG	CTC	TAC	CTG	CTC	ATT	GGG	CTG	CGA	CTG	CGG	CGG	GAG	AGG	CTG	720
221	M	A	I	M	S	V	L	Y	L	L	I	G	L	R	L	R	R	E	R	L	240
TM-12																					
721	CTG	CTC	ATG	CAG	GAG	GCC	AAG	GGC	AGG	GGC	TCT	GCA	GCA	GCC	AGG	TCC	AGA	TAC	ACC	TGC	780
241	L	L	M	Q	E	A	K	G	R	G	S	A	A	A	R	S	R	Y	T	C	260
TM-13																					
781	AGG	CTC	CAG	CAG	CAC	GAT	CGG	GGC	CGG	AGA	CAA	GTG	ACC	AAG	ATG	CTG	TTT	GTC	CTG	GTC	840
261	R	L	Q	Q	H	D	R	G	R	R	Q	V	T	K	M	L	F	V	L	V	280
TM-14																					
841	GTG	GTG	TTT	GGC	ATC	TGC	TGG	GCC	CCG	TTC	CAC	GCC	GAC	CGC	GTC	ATG	TGG	AGC	GTC	GTG	900
281	V	V	F	G	I	C	W	A	P	F	H	A	D	R	V	M	W	S	V	V	300
TM-15																					
901	TCA	CAG	TGG	ACA	GAT	GGC	CTG	CAC	CTG	GCC	TTC	CAG	CAC	GTG	CAC	GTC	ATC	TCC	GGC	ATC	960
301	S	Q	W	T	D	G	L	H	L	A	F	Q	H	V	H	V	I	S	G	I	320
TM-16																					
961	TTC	TTC	TAC	CTG	GGC	TCG	GCG	GCC	AAC	CCC	GTG	CTC	TAT	AGC	CTC	ATG	TCC	AGC	CGC	TTC	1020
321	F	F	Y	L	G	S	A	A	N	P	V	L	Y	S	L	M	S	S	R	F	340
TM-17																					
1021	CGA	GAG	ACC	TTC	CAG	GAG	GCC	CTG	TGC	CTC	GGG	GCC	TGC	TGC	CAT	CGC	CTC	AGA	CCC	CGC	1080
341	R	E	T	F	Q	E	A	L	C	L	G	A	C	C	H	R	L	R	P	R	360
TM-18																					
1081	CAC	AGC	TCC	CAC	AGC	CTC	AGC	AGG	ATG	ACC	ACA	GGC	AGC	ACC	CTG	TGT	GAT	GTG	GGC	TCC	1140
361	H	S	S	H	S	L	S	R	M	T	T	G	S	T	L	C	D	V	G	S	380
TM-19																					
1141	CTG	GGC	AGC	TGG	GTC	CAC	CCC	CTG	GCT	GGG	AAC	GAT	GGC	CCA	GAG	GCG	CAG	CAA	GAG	ACC	1200
381	L	G	S	W	V	H	P	L	A	G	N	D	G	P	E	A	Q	Q	E	T	400
TM-20																					
1201	GAT	CCA	TCC	TGA																	1212
401	D	P	S	*																	404

FIG. 1—Continued

GenBank databases were monitored daily using the Tblastn program (1) with amino acid sequence from the human GHS-R TM domains 6–7 (residues 265–366). A mouse EST (dEST database Accession No. AA562357, deposited August 18, 1997) derived from a T-cell library was identified with a significant homology score. EST 562357 exhibited good sequence identity (63%

DNA, 36% amino acid) to the 3' end of the gene for the human GHS-R. A murine T cell  $\lambda$ XR cDNA library (Stratagene) was screened with the mouse EST 562357 probe (455 bp in length) under high-stringency conditions. Four partial clones were identified after three rounds of screening. Additional clones were isolated from mouse thymus poly(A)<sup>+</sup> RNA via 5' Race Mara-